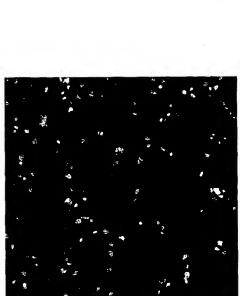
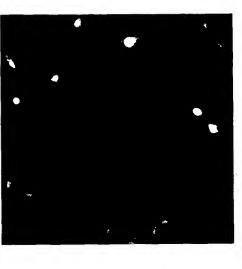
Figure 1



Hoechst 33342 Stain



Green Fluorescent Protein

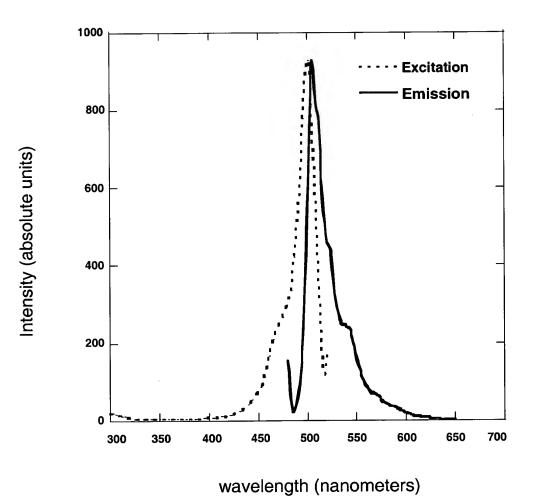
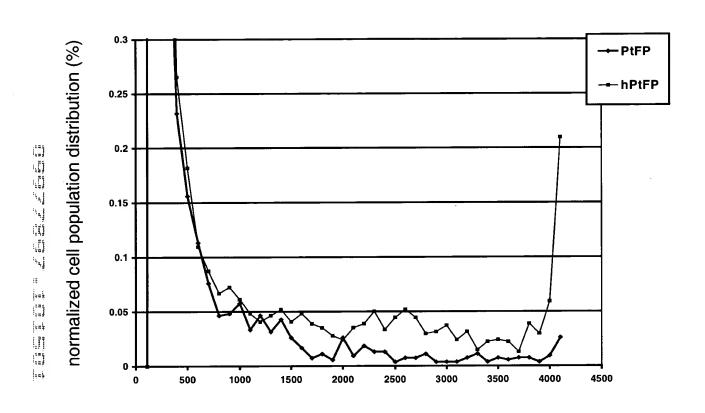


Figure 3

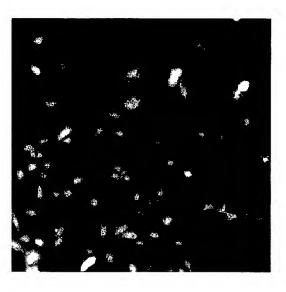


fluorescence intensity

Figure 4



HEK 293 cells



A549 cells

Figure 5

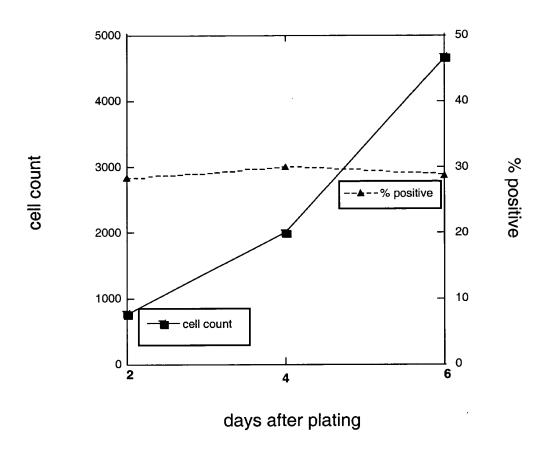


Figure 6

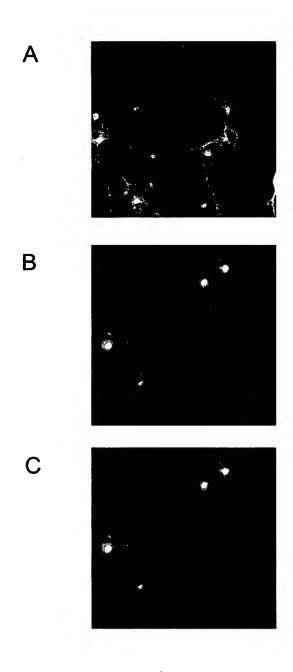


Figure 7

Caspase-3 biosensor

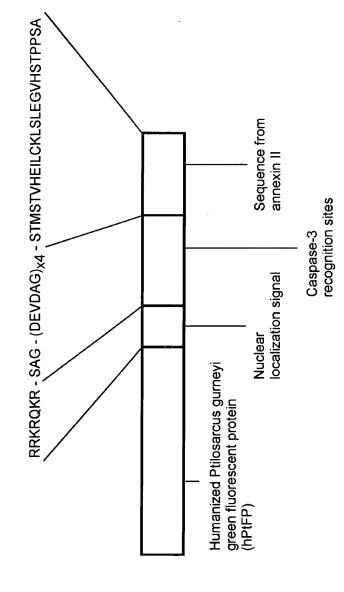
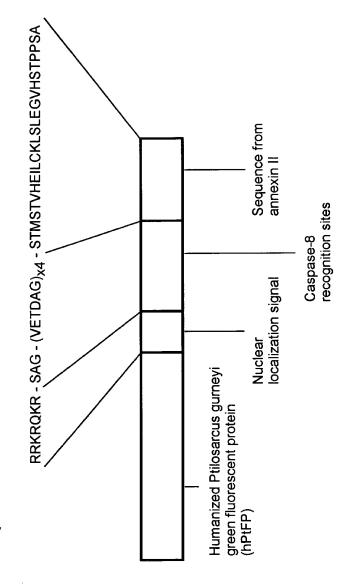


Figure 8





Met

	Met	Val		
±1			Asn	Ara

Asn Arg Asn Val Leu Lys Asn Thr Gly Leu Lys Glu Ile Met Ser Ala Lys Ala ATG AAC CGC AAC GTA TTA AAG AAC ACT GGA CTG AAA GAG ATT ATG TCG GCA AAA GCT PtFP 1 ATG GTG AAC CGG AAC GTG CTG AAG AAC ACC GGC CTG AAG GAG ATC ATG AGC GCC AAG GCC hPtFP \* \* \* +1 Ser Val Glu Gly Ile Val Asn Asn His Val Phe Ser Met Glu Gly Phe Gly Lys Gly Asn 61 AGC GTT GAA GGA ATC GTG AAC AAT CAC GTT TTT TCC ATG GAA GGA TTT GGA AAA GGC AAT PtFP 61 AGC GTG GAG GGC ATC GTG AAC AAC CAC GTG TTC AGC ATG GAG GGC TTC GGC AAG GGC AAC hPtFP +1 Val Leu Phe Gly Asn Gln Leu Met Gln Ile Arg Val Thr Lys Gly Gly Pro Leu Pro Phe 121 GTA TTA TTT GGA AAC CAA TTG ATG CAA ATC CGG GTT ACA AAG GGA GGT CCG TTG CCA TTC PtFP 121 GTG CTG TTC GGC AAC CAG CTG ATG CAG ATC CGG GTG ACC AAG GGC GGC CCT CTG CCC TTC hPtFP +1 Ala Phe Asp Ile Val Ser Ile Ala Phe Gln Tyr Gly Asn Arg Thr Phe Thr Lys Tyr Pro 181 GCT TTC GAT ATT GTT TCC ATA GCT TTC CAA TAC GGG AAT CGC ACT TTC ACG AAA TAC CCA PtFP 181 GCC TTC GAC ATC GTG AGC ATC GCC TTC CAG TAC GGC AAC CGG ACC TTC ACC AAG TAT CCC hPtFP \* \* \* \*\* +1 Asp Asp Ile Ala Asp Tyr Phe Val Gln Ser Phe Pro Ala Gly Phe Phe Tyr Glu Arq Asn PtFP 241 GAC GAC ATT GCG GAC TAC TTT GTT CAA TCA TTC CCG GCT GGA TTT TTC TAC GAA AGA AAT hPtFP 241 GAC GAC ATC GCC GAC TAC TTC GTG CAG AGC TTC CCT GCC GGC TTC TTC TAC GAG CGG AAC \* \*\*\* +1 Leu Arg Phe Glu Asp Gly Ala Ile Val Asp Ile Arg Ser Asp Ile Ser Leu Glu Asp Asp PtFP. 301 CTA CGC TTT GAA GAT GGC GCC ATT GTT GAC ATT CGT TCA GAT ATA AGT TTA GAA GAT GAT hPtPP 301 CTG CGG TTC GAG GAC GGC GCC ATC GTG GAC ATC CGG AGC GAC ATC AGC CTG GAG GAC \* \* \*\*\* \* +1 Lys Phe His Tyr Lys Val Glu Tyr Arg Gly Asn Gly Phe Pro Ser Asn Gly Pro Val Met Ptfg 361 AAG TTC CAC TAC AAA GTG GAG TAT AGA GGC AAC GGT TTC CCT AGT AAC GGA CCC GTG ATG hPtFP 361 AAG TTC CAC TAC AAG GTG GAG TAC CGC GGC AAC GGC TTC CCT AGC AAC GGC CCT GTG ATG \* \* \* 2 = 1 +1 Gln Lys Ala Ile Leu Gly Met Glu Pro Ser Phe Glu Val Val Tyr Met Asn Ser Gly Val 421 CAA AAA GCC ATC CTC GGC ATG GAG CCA TCG TTT GAG GTG GTC TAC ATG AAC AGC GGC GTT PtFP hPtfP 421 CAG AAG GCC ATC CTG GGC ATG GAG CCC AGC TTC GAG GTG GTG TAC ATG AAC AGC GGC GTG \* \*\*\* +1 Leu Val Gly Glu Val Asp Leu Val Tyr Lys Leu Glu Ser Gly Asn Tyr Tyr Ser Cys His 481 CTG GTG GGC GAA GTA GAT CTC GTT TAC AAA CTC GAG TCA GGG AAC TAT TAC TCG TGC CAC hPtFP 481 CTG GTG GGC GAG GTG GAC CTG GTG TAC AAG CTG GAG AGC GGC AAC TAC TAC AGC TGC CAC \*\*\* +1 Met Lys Thr Phe Tyr Arg Ser Lys Gly Gly Val Lys Glu Phe Pro Glu Tyr His Phe Ile 541 ATG AAA ACG TTT TAC AGA TCC AAA GGT GGA GTG AAA GAA TTC CCG GAA TAT CAC TTT ATC PtFP hPtfp 541 ATG AAG ACC TTC TAC CGG AGC AAG GGC GGC GTG AAG GAG TTC CCT GAG TAC CAC TTC ATC \* \* \* +1 His His Arg Leu Glu Lys Thr Tyr Val Glu Glu Gly Ser Phe Val Glu Gln His Glu Thr 601 CAT CAT CGT CTG GAG AAA ACC TAC GTG GAA GGA AGC TTC GTG GAA CAA CAC GAG ACG 601 CAC CAC CGG CTG GAG AAG ACC TAC GTG GAG GAG GGC AGC TTC GTG GAG CAC GAG ACC hPtFP +1 Ala Ile Ala Gln Leu Thr Thr Ile Gly Lys Pro Leu Gly Ser Leu His Glu Trp Val \*\*\*. PLFP 661 GCC ATT GCA CAA CTG ACC ACA ATT GGA AAA CCT CTG GGC TCC CTT CAT GAA TGG GTG TAG hPtfp 661 GCC ATC GCC CAG CTG ACC ACC ATC GGC AAG CCT CTG GGC AGC CTG CAC GAG TGG GTG TAA

## HindIII

- +1 M V N R N V L K N T G

  1 AAG CTT GCC ACC ATG GTG AAC CGG AAC GTG CTG AAG AAC ACC GGC

  TTC GAA CGG TGG TAC CAC TTG GCC TTG CAC GAC TTC TTG TGG CCG
- +1 L K E I M S A K A S V E G I V 46 CTG AAG GAG ATC ATG AGC GCC AAG GCC AGC GTG GAG GGC ATC GTG GAC TTC CTC TAG TAC TCG CGG TTC CGG TCG CAC CTC CCG TAG CAC
- V +1 N N Н V F S Μ Е G F G Κ G 91 AAC AAC CAC GTG TTC AGC ATG GAG GGC TTC GGC AAG GGC AAC GTG TTG TTG GTG CAC AAG TCG TAC CTC CCG AAG CCG TTC CCG TTG CAC
- +1 L F G N Q L M Q I R V T K G G
  136 CTG TTC GGC AAC CAG CTG ATG CAG ATC CGG GTG ACC AAG GGC GGC
  GAC AAG CCG TTG GTC GAC TAC GTC TAG GCC CAC TGG TTC CCG CCG
- +1 P L P F A F D I V S I A F Q Y
  181 CCT CTG CCC TTC GCC TTC GAC ATC GTG AGC ATC GCC TTC CAG TAC
  GGA GAC GGG AAG CGG AAG CTG TAG CAC TCG TAG CGG AAG GTC ATG
- +1 G N R T F T K Y P D D I A D Y
  226 GGC AAC CGG ACC TTC ACC AAG TAT CCC GAC GAC ATC GCC GAC TAC
  CCG TTG GCC TGG AAG TGG TTC ATA GGG CTG CTG TAG CGG CTG ATG
- +1 F V Q S F P A G F F Y E R N L
  271 TTC GTG CAG AGC TTC CCT GCC GGC TTC TTC TAC GAG CGG AAC CTG
  AAG CAC GTC TCG AAG GGA CGG CCG AAG AAG ATG CTC GCC、TTG GAC
- +1 R F E D G A I V D I R S D I S
  316 CGG TTC GAG GAC GGC GCC ATC GTG GAC ATC CGG AGC GAC ATC AGC
  GCC AAG CTC CTG CCG CGG TAG CAC CTG TAG GCC TCG CTG TAG TCG
- +1 L E D D K F H Y K V E Y R G N
  361 CTG GAG GAC GAC AAG TTC CAC TAC AAG GTG GAG TAC CGC GGC AAC
  GAC CTC CTG CTG TTC AAG GTG ATG TTC CAC CTC ATG GCG CCG TTG
- +1 G F P S N G P V M Q K A I L G
  406 GGC TTC CCT AGC AAC GGC CCT GTG ATG CAG AAG GCC ATC CTG GGC
  CCG AAG GGA TCG TTG CCG GGA CAC TAC GTC TTC CGG TAG GAC CCG
- +1 M E P S F E V V Y M N S G V L
  451 ATG GAG CCC AGC TTC GAG GTG GTG TAC ATG AAC AGC GGC GTG CTG
  TAC CTC GGG TCG AAG CTC CAC CAC ATG TAC TTG TCG CCG CAC GAC
- +1 V G E V D L V Y K L E S G N Y
  496 GTG GGC GAG GTG GAC CTG GTG TAC AAG CTG GAG AGC GGC AAC TAC
  CAC CCG CTC CAC CTG GAC CAC ATG TTC GAC CTC TCG CCG TTG ATG
  - +1 Y S C H M K T F Y R S K G G V

- 541 TAC AGC TGC CAC ATG AAG ACC TTC TAC CGG AGC AAG GGC GGC GTG ATG TCG ACG GTG TAC TTC TGG AAG ATG GCC TCG TTC CCG CCG CAC
- +1 K E F P E Y H F I H H R L E K 586 AAG GAG TTC CCT GAG TAC CAC TTC ATC CAC CAC CGG CTG GAG AAG TTC CTC AAG GGA CTC ATG GTG AAG TAG GTG GTG GCC GAC CTC TTC
- +1 T Y V E E G S F V E Q H E T A
  631 ACC TAC GTG GAG GAG GGC AGC TTC GTG GAG CAC GAG ACC GCC
  TGG ATG CAC CTC CTC CCG TCG AAG CAC CTC GTG GTG CTC TGG CGG
- +1 I A Q L T T I G K P L G S L H
  676 ATC GCC CAG CTG ACC ACC ATC GGC AAG CCT CTG GGC AGC CTG CAC
  TAG CGG GTC GAC TGG TGG TAG CCG TTC GGA GAC CCG TCG GAC GTG

## NotI

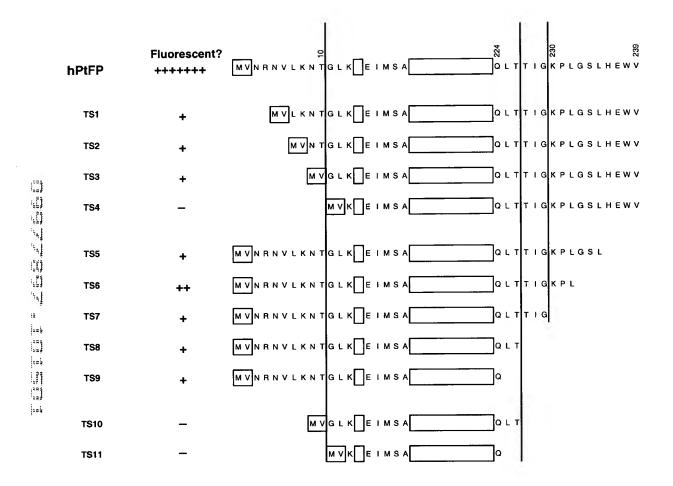
+1 E W V \*
721 GAG TGG GTG TAA AGC GGC CGC
CTC ACC CAC ATT TCG CCG GCG

Figure 10 (continued)

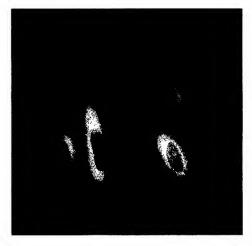
## The coding sequence (from start codon to stop codon):

Figure 11

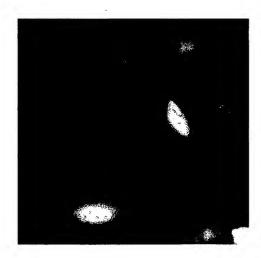
Figure 13



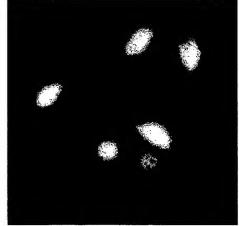
## Figure 14



no treatment



Staurosporine 10 nM 6 hours



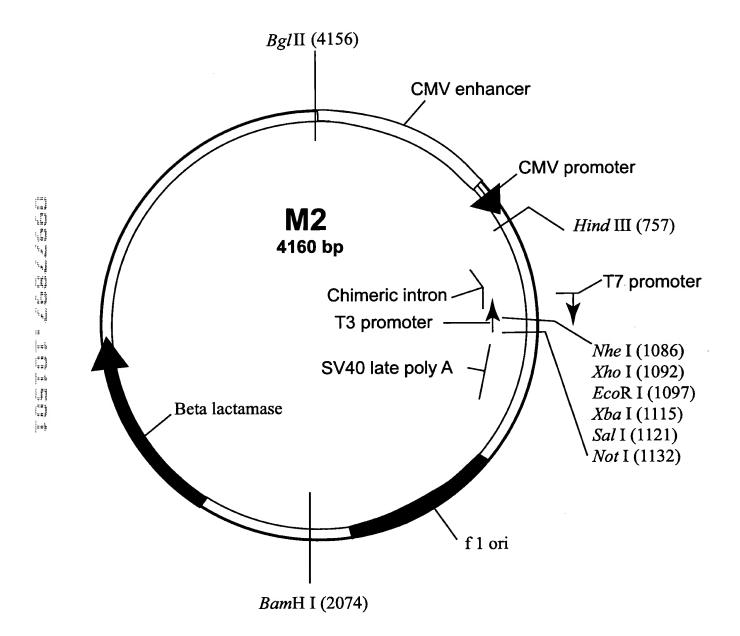
Staurosporine 1 nM 24 hours

```
UUU F 0.45 (185619)
                        UCU S 0.18 (161556)
                                              UAU Y 0.43 (133427)
                                                                    UGU C 0.45 (108740)
  UUC F 0.55 (225633)
                        UCC S 0.22 (192616)
                                              UAC Y 0.57 (174805)
                                                                    UGC C 0.55 (134523)
  UUA L 0.07 ( 79303)
                        UCA S 0.15 (128429)
                                              UAA * 0.29 ( 8187)
                                                                    UGA * 0.50 ( 14381)
  UUG L 0.13 (135218)
                        UCG S 0.06 ( 49456)
                                              UAG * 0.21 (
                                                           5913)
                                                                    UGG W 1.00 (142435)
  CUU L 0.13 (139009)
                        CCU P 0.28 (189374)
                                              CAU H 0.41 (113684)
                                                                    CGU R 0.08 ( 51100)
CUC L 0.20 (210903)
                        CCC P 0.33 (219428)
                                              CAC H 0.59 (162826)
                                                                    CGC R 0.19 (118404)
CUA L 0.07 ( 75667)
                                              CAA Q 0.26 (130857)
                        CCA P 0.27 (182506)
                                                                    CGA R 0.11 ( 68664)
  CUG L 0.40 (435317)
                        CCG P 0.11 ( 76684)
                                              CAG Q 0.74 (377006)
                                                                    CGG R 0.21 (126679)
AUU I 0.35 (174021)
                        ACU T 0.24 (140780)
                                              AAU N 0.46 (186915)
                                                                    AGU S 0.15 (131222)
AUC I 0.49 (240138)
                        ACC T 0.36 (213626)
                                              AAC N 0.54 (218376)
                                                                    AGC S 0.24 (211962)
                                              AAA K 0.42 (262630)
AUA I 0.16 ( 78463)
                        ACA T 0.28 (162837)
                                                                    AGA R 0.20 (125600)
□ AUG M 1.00 (244236)
                        ACG T 0.12 ( 69346)
                                              AAG K 0.58 (359627)
                                                                    AGG R 0.20 (123646)
44
  GUU V 0.18 (119013)
                                              GAU D 0.46 (245435)
                        GCU A 0.26 (202329)
                                                                    GGU G 0.16 (118798)
GUC V 0.24 (160764)
                        GCC A 0.40 (310626)
                                              GAC D 0.54 (287040)
                                                                    GGC G 0.34 (250410)
GUA V 0.11 ( 76398)
GUG V 0.47 (317359)
                        GCA A 0.23 (173010)
                                              GAA E 0.42 (317703)
                                                                    GGA G 0.25 (180955)
                        GCG A 0.11 ( 82647)
                                              GAG E 0.58 (441298)
                                                                    GGG G 0.25 (180001)
[ ]
```

Figure 15

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Figure 16



 ${\tt tcaatattggccattagccatattattcattggttatatagcataaatcaatattggct}$ attggccattgcatacgttgtatctatatcataatatgtacatttatattggctcatgt ccaatatgaccgccatgttggcattgattattgactagttattaatagtaatcaattac ggggtcattagttcatagcccatatatggagttccgcgttacataacttacggtaaatg gcccgcctggctgaccgccaacgacccccgcccattgacgtcaataatgacgtatgtt cccatagtaacgccaatagggactttccattgacgtcaatgggtggagtatttacggta aactgcccacttggcagtacatcaagtgtatcatatgccaagtccgcccctattgacg tcaatgacggtaaatggcccgcctggcattatgcccagtacatgaccttacgggacttt cctacttggcagtacatctacgtattagtcatcgctattaccatggtgatgcggttttg gcagtacaccaatgggcgtggatagcggtttgactcacggggatttccaagtctccacc ccattgacgtcaatgggagtttgttttggcaccaaaatcaacgggactttccaaaatgt cgtaacaactgcgatcgcccgcccgttgacgcaaatgggcggtaggcgtgtacggtgg  ${\tt gaggtctatata} agcagagctcgtttagtgaaccgtcagatcactagaagctttattgc\\$ ggtagtttatcacagttaaattgctaacgcagtcagtgcttctgacacaacagtctcga acttaagctgcagtgactctcttaaggtagccttgcagaagttggtcgtgaggcactgg  $\tt gcaggtaagtatcaaggttacaagacaggtttaaggagaccaatagaaactgggcttgt$ cgagacagagagactcttgcgtttctgataggcacctattggtcttactgacatccac tttgcctttctctccacaggtgtccactcccagttcaattacagctcttaaggctagag  ${\tt tacttaatac} gactcactataggctagcctcgagaattcacgcgtggtacctctagagt$ cgacccgggcggccgcttccctttagtgagggttaatgcttcgagcagacatgataaga tacattgatgagtttggacaaaccacaactagaatgcagtgaaaaaaatgctttatttg tgaaatttgtgatgctattgctttatttgtaaccattataagctgcaataaacaagtta cgtaatagcgaagagcccgcaccgatcgcccttcccaacagttgcgcagcctgaatgg  $\verb|cgaatggacgccctgtagcggcgcattaagcgcgggtgtgtggttacgcgcag|\\$  $\verb|ttctcgccacg| ttcgccggctttccccgtcaagctctaaatcgggggctccctttaggg|$  $\verb|ttccgatttagtgctttacggcacctcgaccccaaaaaacttgattagggtgatggttc|$  ${\tt acgtagtgggccatcgccctgatagacggtttttcgccctttgacgttggagtccacgt}$ tctttaatagtggactcttgttccaaactggaacaacactcaaccctatctcggtctat tcttttgatttataagggattttgccgatttcggcctattggttaaaaaatgagctgat ttaacaaaaatttaacgcgaattttaacaaaatattaacgcttacaatttcctgatgcg gtattttctccttacgcatctgtgcggtatttcacaccgcatacgcggatctgcgcagc  ${\tt accatggcctgaaataacctctgaaagaggaacttggttaggtaccttctgaggcggaa}$ agaaccaggatccgcgtatggtgcactctcagtacaatctgctctgatgccgcatagtt aagccagcccgacacccgccaacacccgctgacgcgccctgacgggcttgtctgctcc cggcatccgcttacagacaagctgtgaccgtctccgggagctgcatgtgtcagaggttt tcaccgtcatcaccgaaacgcgcgagacgaaagggcctcgtgatacgcctatttttata  $\verb|ggtta| at \verb|gata| at \verb|aata| at \verb|ggtttc| tta| acgtca| \verb|ggtga| acttttc| cggggaaatg$ tgcgcggaacccctatttgtttatttttctaaatacattcaaatatgtatccgctcatg agacaataaccctgataaatgcttcaataatattgaaaaaggaagagtatgagtattca acatttccgtgtcgcccttattcccttttttgcggcatttttgccttcctgtttttgctc acccagaaacgctggtgaaagtaaaagatgctgaagatcagttgggtgcacgagtgggt tacatcgaactggatctcaacagcggtaagatccttgagagttttcgccccgaagaacg

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Figure 17 (continued)

Figure 18

